

SEQUENCE LISTING

<110> Dalemans, Wilfried L.J. Gerard, Catherine Marie Ghislaine <120> Compositions Comprising Human Pap

<120> Compositions Comprising Human Papilloma Virus Proteins
 and Fusion Proteins Adjuvanted with a CpG Oligonucleotide
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<140> 09/581,976
<141> 2000-06-20
<150> PCT/EP98/08563

<150> GB 9727262.9 <151> 1997-12-24

<151> 1998-12-18

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<212> PRT

<213> Artificial Sequence

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<223> Chimaeric protein (protein D from Haemophilius influenzae B and E7 from Human papilloma virus type 16)

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Glu Asp Leu Leu Met Gly Thr Leu Gly Ile Val Cys Pro Ile Cys Ser

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            16)
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caaagtttag aaatgacaga aaactttgaa accatggcca tgcatggaga tacacctaca
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ttgcatgaat atatgttaga tttgcaacca gagacaactg atctctactg ttatgagcaa
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ttaaatgaca gctcagagga ggaggatgaa atagatggtc cagctggaca agcagaaccg
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gacagagece attacaatat tgtaacettt tgttgcaagt gtgaetetae getteggttg
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tgcgtacaaa gcacacgt agacattcgt actttggaag acctgttaat gggcacacta
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caaagtttag aaatgacaga aaactttgaa accatggcca tgtttcagga cccacaggag 360
cgacccagaa agttaccaca gttatgcaca gagctgcaaa caactataca tgatataata 420
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ttagaacagc aatacaacaa accgttgtgt gatttgttaa ttaggtgtat taactgtcaa 660
aagccactgt gtcctgaaga aaagcaaaga catctggaca aaaagcaaag attccataat 720
ataaggggtc ggtggaccgg tcgatgtatg tcttgttgca gatcatcaag aacacgtaga 780
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<223> Chimaeric protein (protein D from Haemophilius influenzae B and E6 from Human papilloma virus type 16)

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<223> Chimaeric protein (protein D from Haemophilius influenzae B and E6E7 fusion from Human papilloma virus type 16)

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cgacccagaa agttaccaca gttatgcaca gagctgcaaa caactataca tgatataata
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                                                                       480
                                                                       540
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aagttttatt ctaaaattag tgagtataga cattattgtt atagtttgta tggaacaaca
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ttagaacagc aatacaacaa accgttgtgt gatttgttaa ttaggtgtat taactgtcaa
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aagccactgt gtcctgaaga aaagcaaaga catctggaca aaaagcaaag attccataat
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ataaggggtc ggtggaccgg tcgatgtatg tcttgttgca gatcatcaag aacacgtaga
                                                                       780
                                                                       840
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ccagagacaa ctgatctcta ctgttatgag caattaaatg acagctcaga ggaggaggat
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gaaatagatg gtccagctgg acaagcagaa ccggacagag cccattacaa tattgtaacc
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ttttgttgca agtgtgactc tacgcttcgg ttgtgcgtac aaagcacaca cgtagacatt
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cgtactttgg aagacctgtt aatgggcaca ctaggaattg tgtgccccat ctgttctcag
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<211> 371

<212> PRT

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<223> Chimaeric protein (protein D from Haemophilius influenzae B and E6E7 fusion from Human papilloma virus type 16)

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280
Tyr Glu Gln Leu Asn Asp Ser Ser Glu Glu Glu Asp Glu Ile Asp Gly
                        295
Pro Ala Gly Gln Ala Glu Pro Asp Arg Ala His Tyr Asn Ile Val Thr
                    310
                                        315
Phe Cys Cys Lys Cys Asp Ser Thr Leu Arg Leu Cys Val Gln Ser Thr
                325
                                    330
His Val Asp Ile Arg Thr Leu Glu Asp Leu Leu Met Gly Thr Leu Gly
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Ile Val Cys Pro Ile Cys Ser Gln Lys Pro Thr Ser Gly His His His
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His His His
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     <210> 7
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            influenzae B and mutated E7 from Human papilloma
            virus type 16)
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cttgcgtttg cacaacaggc tgattattta gagcaagatt tagcaatgac taaggatggt
                                                                       180
cgtttagtgg ttattcacga tcacttttta gatggcttga ctgatgttgc gaaaaaattc
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ccacatcgtc atcgtaaaga tggccgttac tatgtcatcg actttacctt aaaagaaatt
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caaagtttag aaatgacaga aaactttgaa accatggcca tgcatggaga tacacctaca
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                                                                       420
ttaaatgaca gctcagagga ggaggatgaa atagatggtc cagctggaca agcagaaccg
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tgcgtacaaa gcacacgt agacattcgt actttggaag acctgttaat gggcacacta
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taa
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            virus type 16)
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Ser Asp Lys Ile Ile Ile Ala His Arg Gly Ala Ser Gly Tyr Leu Pro
Glu His Thr Leu Glu Ser Lys Ala Leu Ala Phe Ala Gln Gln Ala Asp
                            40
Tyr Leu Glu Gln Asp Leu Ala Met Thr Lys Asp Gly Arg Leu Val Val
                        55
                                            60
Ile His Asp His Phe Leu Asp Gly Leu Thr Asp Val Ala Lys Lys Phe
                    70
Pro His Arg His Arg Lys Asp Gly Arg Tyr Tyr Val Ile Asp Phe Thr
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90
Leu Lys Glu Ile Gln Ser Leu Glu Met Thr Glu Asn Phe Glu Thr Met
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            100
Ala Met His Gly Asp Thr Pro Thr Leu His Glu Tyr Met Leu Asp Leu
                            120
Gln Pro Glu Thr Thr Asp Leu Tyr Gly Tyr Gln Gln Leu Asn Asp Ser
                        135
Ser Glu Glu Glu Asp Glu Ile Asp Gly Pro Ala Gly Gln Ala Glu Pro
                    150
                                        155
Asp Arg Ala His Tyr Asn Ile Val Thr Phe Cys Cys Lys Cys Asp Ser
                                    170
Thr Leu Arg Leu Cys Val Gln Ser Thr His Val Asp Ile Arg Thr Leu
                                185
Glu Asp Leu Leu Met Gly Thr Leu Gly Ile Val Cys Pro Ile Cys Ser
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Gln Lys Pro Thr Ser Gly His His His His His
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                        215
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            pneumoniae and E6 from Human papilloma virus type
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cacacagacg gcaactggta ctggttcgac aactcaggcg aaatggctac aggctggaag
                                                                       180
                                                                       240
aaaatcgctg ataagtggta ctatttcaac gaagaaggtg ccatgaagac aggctgggtc
aagtacaagg acacttggta ctacttagac gctaaagaag gcgccatggt atcaaatgcc
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tttatccagt cagcggacgg aacaggctgg tactacctca aaccagacgg aacactggca
                                                                       360
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gacaggccag aattggccag catgctggac atggccatgt ttcaggaccc acaggagcga
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cccagaaagt taccacagtt atgcacagag ctgcaaacaa ctatacatga tataatatta
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gaatgtgtgt actgcaagca acagttactg cgacgtgagg tatatgactt tgcttttcgg
gatttatgca tagtatatag agatgggaat ccatatgctg tatgtgataa atgtttaaag
                                                                       600
ttttattcta aaattagtga gtatagacat tattgttata gtttgtatgg aacaacatta
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gaacagcaat acaacaaacc gttgtgtgat ttgttaatta ggtgtattaa ctgtcaaaag
                                                                       720
                                                                       780
ccactgtgtc ctgaagaaaa gcaaagacat ctggacaaaa agcaaagatt ccataatata
aggggtcggt ggaccggtcg atgtatgtct tgttgcagat catcaagaac acgtagagaa
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            pneumoniae and E6 from Human papilloma virus type
            16)
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Phe Glu Lys Ile Asn Gly Thr Trp Tyr Tyr Phe Asp Ser Ser Gly Tyr
            20
                                25
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Met Leu Ala Asp Arg Trp Arg Lys His Thr Asp Gly Asn Trp Tyr Trp
Phe Asp Asn Ser Gly Glu Met Ala Thr Gly Trp Lys Lys Ile Ala Asp
                        55
Lys Trp Tyr Tyr Phe Asn Glu Glu Gly Ala Met Lys Thr Gly Trp Val
                                         75
                    70
Lys Tyr Lys Asp Thr Trp Tyr Tyr Leu Asp Ala Lys Glu Gly Ala Met
                                     90
                85
Val Ser Asn Ala Phe Ile Gln Ser Ala Asp Gly Thr Gly Trp Tyr Tyr
                                                     110
                                 105
            100
Leu Lys Pro Asp Gly Thr Leu Ala Asp Arg Pro Glu Leu Ala Ser Met
                            120
                                                 125
Leu Asp Met Ala Met Phe Gln Asp Pro Gln Glu Arg Pro Arg Lys Leu
                                             140
                        135
Pro Gln Leu Cys Thr Glu Leu Gln Thr Thr Ile His Asp Ile Ile Leu
                                         155
                    150
Glu Cys Val Tyr Cys Lys Gln Gln Leu Leu Arg Arg Glu Val Tyr Asp
                                                         175
                                     170
                165
Phe Ala Phe Arg Asp Leu Cys Ile Val Tyr Arg Asp Gly Asn Pro Tyr
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                                 185
            180
Ala Val Cys Asp Lys Cys Leu Lys Phe Tyr Ser Lys Ile Ser Glu Tyr
                             200
                                                 205
         195
Arg His Tyr Cys Tyr Ser Leu Tyr Gly Thr Thr Leu Glu Gln Gln Tyr
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                         215
Asn Lys Pro Leu Cys Asp Leu Leu Ile Arg Cys Ile Asn Cys Gln Lys
                                         235
                     230
Pro Leu Cys Pro Glu Glu Lys Gln Arg His Leu Asp Lys Lys Gln Arg
                                     250
                 245
 Phe His Asn Ile Arg Gly Arg Trp Thr Gly Arg Cys Met Ser Cys Cys
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             260
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             pneumoniae and E7 from Human papilloma virus type
             16)
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                                                                         120
                                                                         180
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 aaaatcgctg ataagtggta ctatttcaac gaagaaggtg ccatgaagac aggctgggtc
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 aagtacaagg acacttggta ctacttagac gctaaagaag gcgccatggt atcaaatgcc
                                                                         300
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                                                                         360
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                                                                         420
 catgaatata tgttagattt gcaaccagag acaactgatc tctactgtta tgagcaatta
                                                                         480
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                                                                         540
 agageceatt acaatattgt aacettttgt tgeaagtgtg aetetaeget teggttgtge
                                                                         600
 gtacaaagca cacacgtaga cattcgtact ttggaagacc tgttaatggg cacactagga
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            20
                                25
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                            40
Phe Asp Asn Ser Gly Glu Met Ala Thr Gly Trp Lys Lys Ile Ala Asp
                        55
Lys Trp Tyr Tyr Phe Asn Glu Glu Gly Ala Met Lys Thr Gly Trp Val
                    70
                                        75
Lys Tyr Lys Asp Thr Trp Tyr Tyr Leu Asp Ala Lys Glu Gly Ala Met
                85
                                    90
Val Ser Asn Ala Phe Ile Gln Ser Ala Asp Gly Thr Gly Trp Tyr Tyr
                                105
                                                     110
Leu Lys Pro Asp Gly Thr Leu Ala Asp Arg Pro Glu Leu Ala Ser Met
                            120
                                                 125
Leu Asp Met Ala Met His Gly Asp Thr Pro Thr Leu His Glu Tyr Met
                        135
                                             140
Leu Asp Leu Gln Pro Glu Thr Thr Asp Leu Tyr Cys Tyr Glu Gln Leu
                    150
                                        155
Asn Asp Ser Ser Glu Glu Glu Asp Glu Ile Asp Gly Pro Ala Gly Gln
                                     170
Ala Glu Pro Asp Arg Ala His Tyr Asn Ile Val Thr Phe Cys Cys Lys
            180
                                185
Cys Asp Ser Thr Leu Arg Leu Cys Val Gln Ser Thr His Val Asp Ile
        195
                            200
                                                 205
Arg Thr Leu Glu Asp Leu Leu Met Gly Thr Leu Gly Ile Val Cys Pro
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                                             220
Ile Cys Ser Gln Lys Pro Thr Ser Gly His His His His His
225
                    230
                                        235
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            virus type 16)
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cacacagacg gcaactggta ctggttcgac aactcaggcg aaatggctac aggctggaag
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aaaatcgctg ataagtggta ctatttcaac gaagaaggtg ccatgaagac aggctgggtc
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aagtacaagg acacttggta ctacttagac gctaaagaag gcgccatggt atcaaatgcc
                                                                       300
tttatccagt cagcggacgg aacaggctgg tactacctca aaccagacgg aacactggca
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cccagaaagt taccacagtt atgcacagag ctgcaaacaa ctatacatga tataatatta

480

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ttttattcta aaattagtga gtatagacat tattgttata gtttgtatgg aacaacatta
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                                                                       720
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acccagctga tgcatggaga tacacctaca ttgcatgaat atatgttaga tttgcaacca
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gagacaactg atctctactg ttatgagcaa ttaaatgaca gctcagagga ggaggatgaa
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atagatggtc cagctggaca agcagaaccg gacagagccc attacaatat tgtaaccttt
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tgttgcaagt gtgactctac gcttcggttg tgcgtacaaa gcacacacgt agacattcgt
                                                                      1080
actttggaag acctgttaat gggcacacta ggaattgtgt gccccatctg ttctcagaaa
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<211> 390

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimaeric protein (Clyta from Streptococcus pneumoniae and E6E7 fusion from Human papilloma virus type 16)

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Pro Thr Leu His Glu Tyr Met Leu Asp Leu Gln Pro Glu Thr Thr Asp
                        295
                                             300
Leu Tyr Cys Tyr Glu Gln Leu Asn Asp Ser Ser Glu Glu Glu Asp Glu
                    310
                                         315
Ile Asp Gly Pro Ala Gly Gln Ala Glu Pro Asp Arg Ala His Tyr Asn
                325
                                     330
Ile Val Thr Phe Cys Cys Lys Cys Asp Ser Thr Leu Arg Leu Cys Val
                                345
            340
Gln Ser Thr His Val Asp Ile Arg Thr Leu Glu Asp Leu Leu Met Gly
                            360
                                                 365
Thr Leu Gly Ile Val Cys Pro Ile Cys Ser Gln Lys Pro Thr Ser Gly
                        375
                                             380
His His His His His
385
                    390
      <210> 15
      <211> 684
      <212> DNA
      <213> Artificial Sequence
      <220>
      <223> Chimaeric protein (protein D from Haemophilius
            influenzae B and E7 from Human papilloma virus type
            18)
      <400> 15
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                                                                        60
attattgctc accgtggtgc tagcggttat ttaccagagc atacgttaga atctaaagca
                                                                       120
cttgcgtttg cacaacaggc tgattattta gagcaagatt tagcaatgac taaggatggt
                                                                       180
cgtttagtgg ttattcacga tcacttttta gatggcttga ctgatgttgc gaaaaaattc
                                                                       240
ccacategte ategtaaaga tggeegttae tatgteateg aetttaeett aaaagaaatt
                                                                       300
caaagtttag aaatgacaga aaactttgaa accatggcca tgcatggacc taaggcaaca
                                                                       360
ttgcaagaca ttgtattgca tttagagccc caaaatgaaa ttccggttga ccttctatgt
                                                                       420
cacgagcaat taagcgactc agaggaagaa aacgatgaaa tagatgaagt taatcatcaa
                                                                       480
catttaccag cccgacgagc cgaaccacaa cgtcacacaa tgttgtgtat gtgttgtaag
                                                                       540
tgtgaagcca gaattgagct agtagtagaa agctcagcag acgaccttcg agcattccag
                                                                       600
cagctgtttc tgaacaccct gtcctttgtg tgtccgtggt gtgcatccca gcagactagt
                                                                       660
ggccaccatc accatcacca ttaa
                                                                       684
      <210> 16
      <211> 227
      <212> PRT
      <213> Artificial Sequence
      <220>
      <223> Chimaeric protein (protein D from Haemophilius
            influenzae B and E7 from Human papilloma virus type
            18)
      <400> 16
Met Asp Pro Ser Ser His Ser Ser Asn Met Ala Asn Thr Gln Met Lys
                                     10
Ser Asp Lys Ile Ile Ile Ala His Arg Gly Ala Ser Gly Tyr Leu Pro
            20
                                25
Glu His Thr Leu Glu Ser Lys Ala Leu Ala Phe Ala Gln Gln Ala Asp
Tyr Leu Glu Gln Asp Leu Ala Met Thr Lys Asp Gly Arg Leu Val Val
Ile His Asp His Phe Leu Asp Gly Leu Thr Asp Val Ala Lys Lys Phe
65
                    70
                                         75
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Leu Lys Glu Ile Gln Ser Leu Glu Met Thr Glu Asn Phe Glu Thr Met
                                105
Ala Met His Gly Pro Lys Ala Thr Leu Gln Asp Ile Val Leu His Leu
                            120
Glu Pro Gln Asn Glu Ile Pro Val Asp Leu Leu Cys His Glu Gln Leu
                        135
Ser Asp Ser Glu Glu Glu Asn Asp Glu Ile Asp Glu Val Asn His Gln
                    150
                                        155
His Leu Pro Ala Arg Arg Ala Glu Pro Gln Arg His Thr Met Leu Cys
                                    170
Met Cys Cys Lys Cys Glu Ala Arg Ile Glu Leu Val Val Glu Ser Ser
                                185
                                                     190
Ala Asp Asp Leu Arg Ala Phe Gln Gln Leu Phe Leu Asn Thr Leu Ser
                            200
                                                205
Phe Val Cys Pro Trp Cys Ala Ser Gln Gln Thr Ser Gly His His His
                        215
His His His
225
      <210> 17
      <211> 109
      <212> PRT
      <213> Escherichia coli
      <400> 17
Met Ser Asp Lys Ile Ile His Leu Thr Asp Asp Ser Phe Asp Thr Asp
Val Leu Lys Ala Asp Gly Ala Ile Leu Val Asp Phe Trp Ala Glu Trp
Cys Gly Pro Cys Lys Met Ile Ala Pro Ile Leu Asp Glu Ile Ala Asp
                            40
Glu Tyr Gln Gly Lys Leu Thr Val Ala Lys Leu Asn Ile Asp Gln Asn
                        55
Pro Gly Thr Ala Pro Lys Tyr Gly Ile Arg Gly Ile Pro Thr Leu Leu
                    70
Leu Phe Lys Asn Gly Glu Val Ala Ala Thr Lys Val Gly Ala Leu Ser
                                    90
Lys Gly Gln Leu Lys Glu Phe Leu Asp Ala Asn Leu Ala
            100
      <210> 18
      <211> 684
      <212> DNA
      <213> Artificial Sequence
      <220>
      <223> Chimaeric protein (protein D from Haemophilius
            influenzae B and mutated E7 from Human papilloma
            virus type 18)
      <400> 18
atggatecaa geageeatte ateaaatatg gegaataeee aaatgaaate agaeaaaate
                                                                        60
attattgctc accgtggtgc tagcggttat ttaccagagc atacgttaga atctaaagca
                                                                       120
cttgcgtttg cacaacaggc tgattattta gagcaagatt tagcaatgac taaggatggt
                                                                       180
cgtttagtgg ttattcacga tcacttttta gatggcttga ctgatgttgc gaaaaaattc
                                                                       240
ccacatcgtc atcgtaaaga tggccgttac tatgtcatcg actttacctt aaaagaaatt
                                                                       300
caaagtttag aaatgacaga aaactttgaa accatggcca tgcatggacc taaggcaaca
                                                                       360
ttgcaagaca ttgtattgca tttagagccc caaaatgaaa ttccggttga ccttctaggt
                                                                       420
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Pro His Arg His Arg Lys Asp Gly Arg Tyr Tyr Val Ile Asp Phe Thr

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caccagcaat taagcgactc agaggaagaa aacgatgaaa tagatggagt taatcatcaa 480 catttaccag cccgacgagc cgaaccacaa cgtcacacaa tgttgtgtat gtgttgtaag 540 tgtgaagcca gaattgagct agtagtagaa agctcagcag acgaccttcg agcattccag 600 cagctgtttc tgaacaccct gtcctttgtg tgtccgtggt gtgcatccca gcagactagt 660 ggccaccatc accatcacca ttaa
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<211> 227

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimaeric protein (protein D from Haemophilius influenzae B and mutated E7 from Human papilloma virus type 18)

<400> 19

Met Asp Pro Ser Ser His Ser Ser Asn Met Ala Asn Thr Gln Met Lys 10 Ser Asp Lys Ile Ile Ile Ala His Arg Gly Ala Ser Gly Tyr Leu Pro Glu His Thr Leu Glu Ser Lys Ala Leu Ala Phe Ala Gln Gln Ala Asp Tyr Leu Glu Gln Asp Leu Ala Met Thr Lys Asp Gly Arg Leu Val Val Ile His Asp His Phe Leu Asp Gly Leu Thr Asp Val Ala Lys Lys Phe 70 Pro His Arg His Arg Lys Asp Gly Arg Tyr Tyr Val Ile Asp Phe Thr 90 Leu Lys Glu Ile Gln Ser Leu Glu Met Thr Glu Asn Phe Glu Thr Met 100 105 Ala Met His Gly Pro Lys Ala Thr Leu Gln Asp Ile Val Leu His Leu 120 125 Glu Pro Gln Asn Glu Ile Pro Val Asp Leu Leu Gly His Gln Gln Leu 135 140 Ser Asp Ser Glu Glu Glu Asn Asp Glu Ile Asp Gly Val Asn His Gln 150 155 His Leu Pro Ala Arg Arg Ala Glu Pro Gln Arg His Thr Met Leu Cys 165 170 Met Cys Cys Lys Cys Glu Ala Arg Ile Glu Leu Val Val Glu Ser Ser 180 185 190 Ala Asp Asp Leu Arg Ala Phe Gln Gln Leu Phe Leu Asn Thr Leu Ser 200 205 Phe Val Cys Pro Trp Cys Ala Ser Gln Gln Thr Ser Gly His His His 210 220 215 His His His 225

<210> 20

<211> 837

<212> DNA

<213> Artificial Sequence

<220>

<223> Chimaeric protein (protein D from Haemophilius influenzae virus B and E6 from Human papilloma virus type 18)

<400> 20

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attattgctc accgtggtgc tagcggttat ttaccagagc atacgttaga atctaaagca
                                                                       120
cttgcgtttg cacaacaggc tgattattta gagcaagatt tagcaatgac taaggatggt
                                                                       180
cgtttagtgg ttattcacga tcacttttta gatggcttga ctgatgttgc gaaaaaattc
                                                                       240
ccacatcgtc atcgtaaaga tggccgttac tatgtcatcg actttacctt aaaagaaatt
                                                                       300
caaagtttag aaatgacaga aaactttgaa accatggcgc gctttgagga tccaacacgg
                                                                       360
cgaccctaca agctacctga tctgtgcacg gaactgaaca cttcactgca agacatagaa
                                                                       420
ataacctgtg tatattgcaa gacagtattg gaacttacag aggtatttga atttgcattt
                                                                       480
aaagatttat ttgtggtgta tagagacagt ataccgcatg ctgcatgcca taaatgtata
                                                                       540
gatttttatt ctagaattag agaattaaga cattattcag actctgtgta tggagacaca
                                                                       600
ttggaaaaac taactaacac tgggttatac aatttattaa taaggtgcct gcggtgccag
                                                                       660
aaaccgttga atccagcaga aaaacttaga caccttaatg aaaaacgacg atttcacaac
                                                                       720
atagetggge actatagagg ceagtgeeat tegtgetgea acegageaeg acaggaaega
                                                                       780
ctccaacgac gcagagaaac acaagtaact agtggccacc atcaccatca ccattaa
                                                                       837
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<211> 278

<212> PRT

<213> Artificial Sequence

<220>

<223> Chimaeric protein (protein D from Haemophilius influenzae B and E6 from Human papilloma virus type 18)

<400> 21

Met Asp Pro Ser Ser His Ser Ser Asn Met Ala Asn Thr Gln Met Lys 10 Ser Asp Lys Ile Ile Ile Ala His Arg Gly Ala Ser Gly Tyr Leu Pro 20 Glu His Thr Leu Glu Ser Lys Ala Leu Ala Phe Ala Gln Gln Ala Asp 40 Tyr Leu Glu Gln Asp Leu Ala Met Thr Lys Asp Gly Arg Leu Val Val 55 Ile His Asp His Phe Leu Asp Gly Leu Thr Asp Val Ala Lys Lys Phe 70 Pro His Arg His Arg Lys Asp Gly Arg Tyr Tyr Val Ile Asp Phe Thr 90 Leu Lys Glu Ile Gln Ser Leu Glu Met Thr Glu Asn Phe Glu Thr Met 100 105 110 Ala Arg Phe Glu Asp Pro Thr Arg Arg Pro Tyr Lys Leu Pro Asp Leu 120 125 Cys Thr Glu Leu Asn Thr Ser Leu Gln Asp Ile Glu Ile Thr Cys Val 135 140 Tyr Cys Lys Thr Val Leu Glu Leu Thr Glu Val Phe Glu Phe Ala Phe 150 155 Lys Asp Leu Phe Val Val Tyr Arg Asp Ser Ile Pro His Ala Ala Cys 165 170 His Lys Cys Ile Asp Phe Tyr Ser Arg Ile Arg Glu Leu Arg His Tyr 180 185 Ser Asp Ser Val Tyr Gly Asp Thr Leu Glu Lys Leu Thr Asn Thr Gly 200 205 Leu Tyr Asn Leu Leu Ile Arg Cys Leu Arg Cys Gln Lys Pro Leu Asn 220 215 Pro Ala Glu Lys Leu Arg His Leu Asn Glu Lys Arg Arg Phe His Asn 230 235 Ile Ala Gly His Tyr Arg Gly Gln Cys His Ser Cys Cys Asn Arg Ala 250 245 Arg Gln Glu Arg Leu Gln Arg Arg Arg Glu Thr Gln Val Thr Ser Gly 260 265 270 His His His His His

275 <210> 22 <211> 1152 <212> DNA <213> Artificial Sequence <223> Chimaeric protein (protein D from Haemophilius influenzae B and E6E7 fusion from Human papilloma virus type 18) <400> 22 atggatccaa gcagccattc atcaaatatg gcgaataccc aaatgaaatc agacaaaatc attattgete accgtggtge tageggttat ttaccagage atacgttaga atctaaagea 120 cttgcgtttg cacaacaggc tgattattta gagcaagatt tagcaatgac taaggatggt 180 cgtttagtgg ttattcacga tcacttttta gatggcttga ctgatgttgc gaaaaaattc 240 ccacatcgtc atcgtaaaga tggccgttac tatgtcatcg actttacctt aaaagaaatt 300 360 caaagtttag aaatgacaga aaactttgaa accatggcgc gctttgagga tccaacacgg cgaccctaca agctacctga tctgtgcacg gaactgaaca cttcactgca agacatagaa 420 ataacctgtg tatattgcaa gacagtattg gaacttacag aggtatttga atttgcattt 480 aaagatttat ttgtggtgta tagagacagt ataccgcatg ctgcatgcca taaatgtata 540 gatttttatt ctagaattag agaattaaga cattattcag actctgtgta tggagacaca 600 ttggaaaaac taactaacac tgggttatac aatttattaa taaggtgcct gcggtgccag 660 aaaccgttga atccagcaga aaaacttaga caccttaatg aaaaacgacg atttcacaac 720 atagctgggc actatagagg ccagtgccat tcgtgctgca accgagcacg acaggaacga 780 ctccaacgac gcagagaaac acaagtaatg catggaccta aggcaacatt gcaagacatt 840 gtattgcatt tagagcccca aaatgaaatt ccggttgacc ttctatgtca cgagcaatta 900 agcgactcag aggaagaaaa cgatgaaata gatggagtta atcatcaaca tttaccagcc 960 1020 cgacgagccg aaccacaacg tcacacaatg ttgtgtatgt gttgtaagtg tgaagccaga attgagctag tagtagaaag ctcagcagac gaccttcgag cattccagca gctgtttctg 1080 aacaccctgt cetttgtgtg teegtggtgt geateceage agactagtgg ceaccateae 1140 1152 catcaccatt aa <210> 23 <211> 383 <212> PRT <213> Artificial Sequence <220> <223> Chimaeric protein (protein D from Haemophilius influenzae B and E6E7 fusion from Human papilloma virus type 18) <400> 23 Met Asp Pro Ser Ser His Ser Ser Asn Met Ala Asn Thr Gln Met Lys 10 Ser Asp Lys Ile Ile Ile Ala His Arg Gly Ala Ser Gly Tyr Leu Pro 25 Glu His Thr Leu Glu Ser Lys Ala Leu Ala Phe Ala Gln Gln Ala Asp 40 Tyr Leu Glu Gln Asp Leu Ala Met Thr Lys Asp Gly Arg Leu Val Val 55 Ile His Asp His Phe Leu Asp Gly Leu Thr Asp Val Ala Lys Lys Phe

70

Pro His Arg His Arg Lys Asp Gly Arg Tyr Tyr Val Ile Asp Phe Thr

Leu Lys Glu Ile Gln Ser Leu Glu Met Thr Glu Asn Phe Glu Thr Met 105 Ala Arg Phe Glu Asp Pro Thr Arg Arg Pro Tyr Lys Leu Pro Asp Leu

90

60

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Cys Thr Glu Leu Asn Thr Ser Leu Gln Asp Ile Glu Ile Thr Cys Val
                135
Tyr Cys Lys Thr Val Leu Glu Leu Thr Glu Val Phe Glu Phe Ala Phe
                   150
                                       155
Lys Asp Leu Phe Val Val Tyr Arg Asp Ser Ile Pro His Ala Ala Cys
                                    170
His Lys Cys Ile Asp Phe Tyr Ser Arg Ile Arg Glu Leu Arg His Tyr
           180
                               185
Ser Asp Ser Val Tyr Gly Asp Thr Leu Glu Lys Leu Thr Asn Thr Gly
                           200
Leu Tyr Asn Leu Leu Ile Arg Cys Leu Arg Cys Gln Lys Pro Leu Asn
                       215
Pro Ala Glu Lys Leu Arg His Leu Asn Glu Lys Arg Arg Phe His Asn
                   230
                                        235
Ile Ala Gly His Tyr Arg Gly Gln Cys His Ser Cys Cys Asn Arg Ala
                                   250
Arg Gln Glu Arg Leu Gln Arg Arg Glu Thr Gln Val Met His Gly
           260
                               265
Pro Lys Ala Thr Leu Gln Asp Ile Val Leu His Leu Glu Pro Gln Asn
                           280
Glu Ile Pro Val Asp Leu Cys His Glu Gln Leu Ser Asp Ser Glu
                       295
Glu Glu Asn Asp Glu Ile Asp Gly Val Asn His Gln His Leu Pro Ala
                   310
                                        315
Arg Arg Ala Glu Pro Gln Arg His Thr Met Leu Cys Met Cys Cys Lys
               325
                                    330
Cys Glu Ala Arg Ile Glu Leu Val Val Glu Ser Ser Ala Asp Asp Leu
           340
                               345
                                                    350
Arg Ala Phe Gln Gln Leu Phe Leu Asn Thr Leu Ser Phe Val Cys Pro
       355
                           360
                                                365
Trp Cys Ala Ser Gln Gln Thr Ser Gly His His His His His
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                       375
                                            380
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      <212> DNA
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     <223> Synthetic
     <400> 25
tctcccagcg tgcgccat
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     <210> 26
      <211> 30
      <212> DNA
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120

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       <223> Synthetic
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       <211> 6
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       <223> Synthetic
      <400> 27
                                                                           6
rrcgyy
       <210> 28
       <211> 9
       <212> PRT
       <213> Artificial Sequence
       <220>
       <223> E.coli
      <400> 28
Thr Ser Gly His His His His His
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